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IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant(s) : Lamb et al.  
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Corresponding Int'l. App. : PCT/GB97/03058  
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THOMAS J. KOWALSKI, Reg. No. 32,147

Name of Applicant, Assignee or Registered Representative

*Thomas J. Kowalski*  
Signature

18 Apr 02  
Date of Signature

COMMUNICATION

Assistant Commissioner for Patents  
Washington, D.C. 20231

Sir:

Pursuant to the provisions of 37 C.F.R. 1.28(b), Applicants wish to notify the U.S. Patent  
and Trademark Office that they are no longer entitled to small entity status under 37 C.F.R. 1.9(f).

Accordingly, all fees should be calculated on the basis of large entity status.

Please charge any additional fees required to Deposit Account No. 50-0320.

Respectfully submitted,

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**Notch Ligands Known in the Art Before the Filing Date**

<b>Ligand</b>	<b>Publication</b>	<b>Date</b>
Drosophila Delta	<b>Kopczynski et al.</b> , 1988, Genes Dev.2, 1723-1735: "Delta, a Drosophila neurogenic gene, is transcriptionally complex and encodes a protein related to blood coagulation factors and epidermal growth factor of vertebrates"	1988
Drosophila Serrate	<b>Fleming et al.</b> , 1990, Genes Dev. 4, 2188-2201: "The gene Serrate encodes a putative EGF-like transmembrane protein essential for proper ectodermal development in Drosophila melangaster"	1990
C. elegans Lag-2	<b>Tax et al.</b> , 1994, Nature 368, 150-154: "Sequence of C. elegans lag-2 reveals a cell-signalling domain shared with Delta and Serrate of Drosophila"	1994
Rat Jagged 1	<b>Lindsell et al.</b> , 1995, Cell 80, 909-917: "Jagged: A Mammalian Ligand That Activates Notch 1"	March, 1995
Mouse Delta 1	<b>Bettenhasen et al.</b> , 1995, Development 121, 2407-2418: "Transient and restricted expression during mouse embryogenesis of Dll 1, a murine gene closely related to Drosophila Delta"	1995
Chick and Xenopus Delta 1	<b>Henrique et al.</b> , 1995, Nature 375, 787-790: "Expression of a Delta homologue in prospective neurons in the chick"	June, 1995
Chick Serrate 1	<b>Myat et al.</b> , 1996, Dev. Biol. 174, 233-247: "A chick homologue of Serrate and its relationship with Notch and Delta homologues during central neurogenesis"	1996
Chick Serrate 2	<b>Hayashi et al.</b> , 1996, Int. J. Dev. Biol. 40, 1089-1096: "Isolation of a novel chick homolog of Serrate and its coexpression with C-Notch-1 in chick development"	December, 1996
Rat Jagged 2	<b>Shawber et al.</b> , 1996, Dev. Biol. 180, 370-376: "Jagged 2: a Serrate-like gene expressed during rat embryogenesis"	November, 1996
Human Jagged 1	<b>Ish-Horowicz et al.</b> , WO 96/27610 "Nucleotide and Protein Sequences of Vertebrate Serrate Genes and Methods Based Thereon"	September, 1996
Zebra Fish Delta 1	<b>Dornseifer et al.</b> , 1997, Mech. Dev. 63, 159-171: "Overexpression of a zebrafish homologue of the Drosophila neurogenic gene Delta perturbs differentiation of primary neurons and somite development"	May, 1997

Human Jagged 1	<b>Oda et al.</b> , 1997, Genomics 43, 376-379: "Identification and cloning of the human homolog (JAG1) of the rat Jagged 1 gene from the Alagille syndrome critical region at 20p12"	August, 1997
Human Jagged 2	<b>Luo et al.</b> , 1997, Mol. Cell Biol. 17, 6057-6067: "Isolation and functional analysis of a cDNA for human Jagged 2, a gene encoding a ligand for the Notch 1 receptor" (also reports sequences for rat Jagged 1 and Jagged 2)	October, 1997
Human Delta 1	<b>Mann et al.</b> , 1997, GenBank AF003522; Direct Submission	May, 1997
Human Delta 1	<b>Ish-Horowicz et al.</b> , WO 97/01571, "Nucleotide and Protein Sequences of Vertebrate Delta Genes and Methods Based Thereon"	January, 1997